FILE 'HOME' ENTERED AT 12:18:16 ON 01 DEC 2006

=> file registry

=>

Uploading C:\Program Files\Stnexp\Queries\10665079.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 17 18 19 20 21

ring nodes : 1 2 3 4 5 6

chain bonds :

5-7 7-8 7-11 8-9 8-10 9-12 12-13 13-14 14-15 15-16 16-17 17-18 17-20

18-19 18-21 ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :

5-7 7-8 8-10 17-18 17-20 18-19

exact bonds :

7-11 8-9 9-12 12-13 13-14 14-15 15-16 16-17 18-21

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6

isolated ring systems :

containing 1 :

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS

11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS

19:CLASS 20:CLASS

21:CLASS

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR

Structure attributes must be viewed using STN Express query preparation.

=> s 11

SAMPLE SEARCH INITIATED 12:18:44 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 175 TO ITERATE

100.0% PROCESSED 175 ITERATIONS

5 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

2707 TO 4293

PROJECTED ANSWERS:

5 TO 234

5 SEA SSS SAM L1

=> s l1 exa full

FULL SEARCH INITIATED 12:18:50 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 48 TO ITERATE

100.0% PROCESSED

48 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

L32 SEA EXA FUL L1

=> d scan

2 ANSWERS REGISTRY COPYRIGHT 2006 ACS on STN L3

IN Octanediamide, N-hydroxy-N'-(phenyl-4-t)- (9CI)

MF C14 H19 N2 O3 T

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):1

L3 2 ANSWERS REGISTRY COPYRIGHT 2006 ACS on STN

IN Octanediamide, N-hydroxy-N'-phenyl- (9CI)

MF C14 H20 N2 O3

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

=> file medline, caplus, uspatfull, wpids

=> s 13

SAMPLE SEARCH INITIATED 12:19:26 FILE 'WPIDS'

SAMPLE SCREEN SEARCH COMPLETED - 15 TO ITERATE

100.0% PROCESSED 15 IT

15 ITERATIONS

4 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 34 TO 266
PROJECTED ANSWERS: 4 TO 99

L4 378 L3

=> s 14 and "b-cell lymphoma"

L5 10 L4 AND "B-CELL LYMPHOMA"

=> d 15 1-10 ibib, abs, hitstr

L5 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2006:909614 CAPLUS Full-text

DOCUMENT NUMBER:

145:363490

TITLE:

The medicinal composite for treating B

cell lymphoma

INVENTOR(S):

Chen, Zhu; Chen, Saijuan; Zhao, Weili; Wang, Lan; Liu,

Yuanhua; Yan, Jinsong

PATENT ASSIGNEE(S):

Ruijin Hospital Affiliated To Shanghai Jiaotong University School of Medicine, Peop. Rep. China

SOURCE:

Faming Zhuanli Shenqing Gongkai Shuomingshu, 20pp.

CODEN: CNXXEV

DOCUMENT TYPE:

Patent

LANGUAGE:

Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KI		DATE	APPLICATION NO.	DATE		
CN 1824307	Α	20060830	CN 2005-10112147	20051228		
PRIORITY APPLN. INFO.:			CN 2005-10112147	20051228		
AD This invention rel	12+05 +0	. a medicinal	composite for treating	B cell lym		

AB This invention relates to a medicinal composite for treating B cell lymphoma, which comprises Rituximab 1-100 μ g/mL, SAHA (suberoyl anilide hydroxamic acid) 1-10 μ M. Title has obvious synergistic action of inhibition of B lymphoma cell growth and breeding, synergistic inducement of apoptosis, synergistic inhibition of NF- κ B activity and lowing BCL-XL expression.

IT 149647-78-9, Suberoyl anilide hydroxamic acid

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(medicinal composite for treating B cell lymphadenopathy)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

L5 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:459141 CAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 144:460489

TITLE: Histone Deacetylase Inhibitors Suppress the

Inducibility of Nuclear Factor- κB by Tumor Necrosis Factor- α Receptor-1 Down-regulation Imre, Gabriele; Gekeler, Volker; Leja, Astrid;

AUTHOR(S): Imre, Gabriele; Gekeler, Volker; Leja, Ast

Beckers, Thomas; Boehm, Markus

CORPORATE SOURCE: Therapeutic Area Oncology, ALTANA Pharma AG, Konstanz,

D-78367, Germany

SOURCE: Cancer Research (2006), 66(10), 5409-5418

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER: American Association for Cancer Research

DOCUMENT TYPE: Journal LANGUAGE: English

Recently, the inhibition of histone deacetylase (HDAC) enzymes has attracted AB attention in the oncol. community as a new therapeutic opportunity for hematol. and solid tumors including non-small cell lung cancer (NSCLC). hematol. malignancies, such as diffuse large B -cell lymphoma, the HDAC inhibitor (HDI), suberoylanilide hydroxamic acid (SAHA), has recently entered phase II and III clin. trials. To further advance our understanding of their action on tumor cells, we investigated the possible effect of HDI treatment on the functionality of the nuclear factor-KB (NF-KB) pathway in NSCLC. We found that in the NSCLC cell lines, A549 and NCI-H460, the NF-KB pathway was strongly inducible, for example, by stimulation with tumor necrosis factor- α (TNF-α). Incubation of several NSCLC cell lines with HDIs resulted in greatly reduced gene expression of TNF- α receptor-1. HDI-treated A549 and NCI-H460 cells down-regulated TNF- α receptor-1 mRNA and protein levels as well as surface exposure, and consequently responded to $TNF-\alpha$ treatment with reduced IKK phosphorylation and activation, delayed $I\kappa B-\alpha$ phosphorylation, and attenuated NF-kB nuclear translocation and DNA binding. Accordingly, stimulation of NF-kB target gene expression by TNF- α was strongly decreased. In addition, we observed that SAHA displayed antitumor efficacy in vivo against A549 xenografts grown on nude mice. HDIs, therefore, might beneficially contribute to tumor treatment, possibly by reducing the responsiveness of tumor cells to the TNF- α -mediated activation of the NF- κB pathway. These findings also hint at a possible use of HDIs in inflammatory diseases, which are associated with the overprodn. of $TNF-\alpha$, such as rheumatoid arthritis or Crohn's disease.

IT 149647-78-9, Suberoylanilide hydroxamic acid

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(histone deacetylase inhibitors suppress the inducibility of nuclear factor- κB by tumor necrosis factor- α receptor-1 down-regulation)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2006:82439 CAPLUS Full-text

DOCUMENT NUMBER:

144:403895

TITLE:

Clinical experience with intravenous and oral

formulations of the novel histone deacetylase

inhibitor suberoylanilide hydroxamic acid in patients

with advanced hematologic malignancies

AUTHOR (S):

O'Connor, Owen A.; Heaney, Mark L.; Schwartz, Lawrence; Richardson, Stacie; Willim, Robert;

MacGregor-Cortelli, Barbara; Curly, Tracey; Moskowitz, Craig; Portlock, Carol; Horwitz, Steven; Zelenetz, Andrew D.; Frankel, Stanley; Richon, Victoria; Marks,

Paul; Kelly, William K.

CORPORATE SOURCE:

Department of Medicine, Division of Hematologic Oncology, Lymphoma Service, Leukemia Service, Division of Solid Tumor Oncology, Developmental Chemotherapy Service, Genitourinary Oncology Service, Department of Radiology, Department of Nursing, Cell Biology Program, Mem. Sloan-Kettering Cancer Center, Sloan-Kettering Institute, New York, NY, USA

SOURCE:

Journal of Clinical Oncology (2006), 24(1), 166-173

CODEN: JCONDN; ISSN: 0732-183X

PUBLISHER:

American Society of Clinical Oncology

DOCUMENT TYPE: LANGUAGE: Journal English

Purpose: To document the toxicity and activity of the histone deacetylase AB inhibitor suberoylanilide hydroxamic acid (SAHA) in patients with pretreated hematol. malignancies. Patients and Methods: Two formulations of SAHA (i.v. [IV] and oral) have been assessed in two consecutive phase I trials. In both trials, dose escalation was performed in parallel and independently in patients with solid tumors and hematol. malignancies. Eligible patients were required to have adequate hepatic and renal function, an absolute neutrophil count ≥ 500/µL and a platelet count more than 25,000/mL. All patients provided informed consent for study inclusion. Results: A total of 39 patients with hematol. malignancy were enrolled (14 on IV SAHA and 25 on oral SAHA), of whom 35 were treated. The spectrum of diseases included patients with diffuse large B- cell lymphoma (n = 12), Hodgkin's disease (HD; n = 12), multiple myeloma (n = 2), T-cell lymphoma (n = 3), mantle cell lymphoma (n = 3)2), small lymphocytic lymphoma (n = 2), and myeloid leukemia (n = 2). Major adverse events with the oral formulation included fatigue, diarrhea, anorexia, and dehydration, whereas myelosuppression and thrombocytopenia were more prominent with the IV formulation. Typically, the hematol. toxicities resolved shortly after SAHA was stopped. There was no neutropenic fever or neutropenic sepsis. Reduction in measurable tumor was observed in five patients. One patient with transformed small lymphocytic lymphoma met criteria for complete response, whereas another met the criteria for partial response (PR). One patient with refractory HD had a PR, whereas three patients had stable disease for up to 9 mo. Conclusion: These results suggest that SAHA has activity in hematol. malignancies including HD and select subtypes of non-Hodgkin's lymphoma.

149647-78-9, Suberoylanilide hydroxamic acid

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(oral HDAC inhibitor SAHA showed clin. activity, with toxicities like fatigue, diarrhea, anorexia, dehydration, while IV formulation showed myelosuppression, fatigue, infection, thrombocytopenia in hematol. malignancies patient)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

O O | PhNH_C_(CH₂)6_C_NH_OH

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:177856 CAPLUS Full-text

DOCUMENT NUMBER:

142:254579

TITLE:

Method of treating cancer with histone deacetylase

(HDAC) inhibitors

INVENTOR(S):

Bacopoulos, Nicholas G.; Chiao, Judy H.; Miller,

Thomas A.; Paradise, Carolyn M.; Richon, Victoria M.

PATENT ASSIGNEE(S):

Aton Pharma, Inc., USA; Sloan-Kettering Institute for

Cancer Research

SOURCE:

PCT Int. Appl., 107 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 7

PATENT INFORMATION:

PA	rent :	NO.			KIN	D	DATE			APPL	ICAT	ION I	NO.		D	ATE	
	2005				A2 A3		2005		,	WO 2	004-1	US27	943		20	0040	826
	W: RW:	CN, GE, LK, NO, TJ, BW, AZ, EE,	CO, GH, LR, NZ, TM, GH, BY, ES,	CR, GM, LS, OM, TN, GM, KG, FI,	CU, HR, LT, PG, TR, KE, KZ,	CZ, HU, LU, PH, TT, LS, MD, GB,	AU, DE, ID, LV, PL, TZ, MW, RU, GR, CF,	DK, IL, MA, PT, UA, MZ, TJ,	DM, IN, MD, RO, UG, NA, TM, IE,	DZ, IS, MG, RU, UZ, SD, AT, IT,	EC, JP, MK, SC, VC, SL, BE, LU,	EE, KE, MN, SD, VN; SZ, BG, MC,	EG, KG, MW, SE, YU, TZ, CH, NL,	ES, KP, MX, SG, ZA, UG, CY, PL,	FI, KR, MZ, SK, ZM, CM, CZ, PT,	GB, KZ, NA, SL, ZW, ZW, DE, RO,	GD, LC, NI, SY, AM, DK, SE,
		SN,	TD,	TG	·	·			-					J.,			
US	2004	0876	31		A1		2004	0506	Ī	US 2	003-	6500	25		_ 20	0030	826
US	2004	1275	23		A1		2004	0701	•	US 2	003-	6650	79		20	0030	916
ΑU	2004	2661	69		A1		2005	0303		AU 2	004-	2661	69		20	040	826
ΑU	2004	2661	69		A2		2005	0303									
CA	2535	806			AA		2005	0303		CA 2	004-	2535	806		20	040	826
EP	1663	194			A2		2006	0607		EP 2	004-	7824	25		20	040	826
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		IE,	SI,	LT,	LV,	FI,	RO,	CY,	TR,	BG,	CZ,	EE,	HU,	PL,	SK		
NO	2006	0013	48		A		2006	0523	Ì	NO 2	006-	1348	-		20	0060	324

PRIORITY APPLN. INFO.:

US 2003-650025 A1 20030826 US 2003-655079 A 20030916 US 2003-665079 A1 20030916 US 2002-361759P P 20020304 US 2003-379149 A2 20030304 WO 2004-US27943 W 20040826

OTHER SOURCE(S): MARPAT 142:254579

The invention discloses methods for treating cancers, e.g. mesothelioma or lymphoma. More specifically, the invention discloses methods for treating mesothelioma or diffuse large B-cell lymphoma (DLBCL), by administration of pharmaceutical compns. comprising HDAC inhibitors, e.g., suberoylanilide hydroxamic acid (SAHA; preparation described). The oral formulations of the pharmaceutical compns. have favorable pharmacokinetic profiles such as high bioavailability and surprisingly give rise to high blood levels of the active compds. over an extended period of time. The invention further provides a safe, daily dosing regimen of these pharmaceutical compns., which is easy to follow, and which results in a therapeutically effective amount of the HDAC inhibitors in vivo.

IT 149647-78-9P, Saha

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(histone deacetylase inhibitors for cancer treatment)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

PhnH_C_(CH₂)6_C_NH_OH

L5 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:69409 CAPLUS Full-text

DOCUMENT NUMBER: 142:385256

TITLE: Histone deacetylase inhibitors profoundly decrease

proliferation of human lymphoid cancer cell lines

AUTHOR(S): Sakajiri, Sakura; Kumagai, Takashi; Kawamata,

Norihiko; Saitoh, Takayuki; Said, Jonathan W.;

Koeffler, H. Phillip

CORPORATE SOURCE: Division of Hematology/Oncology, Cedars Sinai Medical

Center, UCLA School of Medicine, Los Angeles, CA, USA

SOURCE: Experimental Hematology (New York, NY, United States)

(2005), 33(1), 53-61

CODEN: EXHMA6; ISSN: 0301-472X

PUBLISHER: Elsevier Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

Methylation of tumor suppressor genes is frequently observed in human cancers. These genes are silenced by histone deacetylase (HDAC) recruited by methylated DNA in their promoter regions. HDAC removes acetyl groups from histones and prevents the basic transcriptional machinery access to the target gene, leading to transcriptional repression. HDAC inhibitors (HDACIs) can restore the expression of the tumor suppressor and/or cell cycle regulatory genes in cancer cells and block the cellular proliferation of these cells. In this study, we investigated the in vitro antiproliferative activities of the HDACIs, suberoylanilide hydroxamic acid (SAHA), and valproic acid against 14

human lymphoid cancer cell lines. All of these cell lines were sensitive to the antiproliferative effects of the HDACI. SAHA induced either G1 or G2-M arrest as well as apoptosis. SAHA downregulated cyclin D1 and D2, and upregulated p53, p21, and p27. Chromatin immunopptn. anal. revealed a remarkable increase in the level of acetylated histones associated with the p21 promoter after SAHA treatment. In nude mice, SAHA significantly inhibited growth of a mantle cell lymphoma without major toxic side effects. In summary, HDACIs are promising therapeutic agents for human lymphoid cancers. 149647-78-9, Suberoylanilide hydroxamic acid

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(SAHA showed antiproliferative effect, induced cell cycle arrest, apoptosis, reduced cycline D1, D2, raised p53, p21, p27 in human lymphoid cancer cell lines, showed antitumor activity in NCEBI human mantle lymphoma cell transplanted mouse)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:533980 CAPLUS Full-text

DOCUMENT NUMBER:

141:65091

TITLE:

IT

Methods of treating cancer with histone deacetylase

(HDAC) inhibitors

INVENTOR(S):

Bacopoulos, Nicholas G.; Chiao, Judy H.; Miller,

Thomas A.; Paradise, Carolyn M.; Richon, Victoria M.

PATENT ASSIGNEE(S):

USA

SOURCE:

U.S. Pat. Appl. Publ., 54 pp., Cont.-in-part of U.S.

Ser. No. 379,149.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

Endita

PATENT INFORMATION:

PAT	rent :	NO.			KIN	D :	DATE			APPL	ICAT	ION I	NO.		. D	ATE	
US	2004	 1275:	23		A1	-	2004	0701		US 2	 003-	 6650'	79		2	0030	916
US	2004	0727	35		A1			20040415			US 2003-379149				20030304		
AU	2004	2661	69		A1		2005	0303		AU 2	004-	2661	69		2	0040	826
ΑU	2004	2661	69		A2		2005	0303				•					
CA	2535	806			AA		2005	0303.		CA 2	004-	2535	806		2	0040	826
WO	WO 2005018578 A2				20050303 WO 2004-US27943					20040826							
WO	2005	0185	78		A3		2005	0512									
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
		ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	UZ,	VC,	VN,	YU,	ZA,	ZM,	zw	
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,

AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20060607 EP 2004-782425 EP 1663194 A2 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK 20060523 NO 2006-1348 NO 2006001348 Α 20060324 US 2002-361759P P 20020304 US 2003-379149 A2 20030304 PRIORITY APPLN. INFO.: A 20030826 US 2003-650025 A 20030916 US 2003-655079 US 2003-665079 A 20030916

OTHER SOURCE(S): MARPAT 141:65091

The invention relates to methods of treating cancers, e.g., lymphoma. More specifically, the present invention relates to methods of treating diffuse large B-cell lymphoma (DLBCL), by administration of pharmaceutical compns. comprising HDAC inhibitors, e.g., suberoylanilide hydroxamic acid (preparation described). The oral formulations of the pharmaceutical compns. have favorable pharmacokinetic profiles such as high bioavailability and surprisingly give rise to high blood levels of the active compds. over an extended period of time. The present invention further provides a safe, daily dosing regimen of these pharmaceutical compns., which is easy to follow, and which results in a therapeutically effective amount of the HDAC inhibitors in vivo.

WO 2004-US27943

W 20040826

IT 149647-78-9P, SAHA

RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(histone deacetylase inhibitors for treatment of cancer)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

L5 ANSWER 7 OF 10 USPATFULL on STN

ACCESSION NUMBER: 2006:68089 USPATFULL Full-text

TITLE: Combinations for the treatment of diseases involving

cell proliferation

INVENTOR(S): Munzert, Gerd, Ulm, GERMANY, FEDERAL REPUBLIC OF

Steegmaier, Martin, Wien, AUSTRIA

Baum, Anke, Vienna, AUSTRIA

PATENT ASSIGNEE(S): Boehringer Ingelheim International GmbH, Ingelheim,

GERMANY, FEDERAL REPUBLIC OF (non-U.S. corporation)

NUMBER DATE

PRIORITY INFORMATION: EP 2004-19361 20040814

EP 2004-19448 20040817

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: MICHAEL P. MORRIS, BOEHRINGER INGELHEIM CORPORATION,

900 RIDGEBURY ROAD, P. O. BOX 368, RIDGEFIELD, CT,

06877-0368, US

NUMBER OF CLAIMS: 22 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Page(s)

LINE COUNT: 3176

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Disclosed are pharmaceutical compositions for the treatment of diseases which involve cell proliferation. Also disclosed are methods for the treatment of said diseases, comprising co-administration of a compound 1 of Formula (I) ##STR1## wherein the groups L, R.sup.1, R.sup.2, R.sup.3, R.sup.4 and R.sup.5 have the meanings given herein and of an effective amount of an active compound 2 and/or co-treatment with radiation therapy, in a ratio which provides an additive and synergistic effect, and to the combined use of a compound 1 of Formula (I) and of an effective amount of an active compound 2 and/or radiotherapy for the manufacture of corresponding pharmaceutical combination preparations.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9, SAHA

(preparation of aminopteridinones for use in combination therapy for treatment of cell proliferative diseases)

RN 149647-78-9 USPATFULL

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

L5 ANSWER 8 OF 10 USPATFULL on STN

ACCESSION NUMBER: 2005:331242 USPATFULL <u>Full-text</u>

TITLE: Use of thioredoxin measurements for diagnostics and

treatments

INVENTOR(S): Marks, Paul A., Washington, CT, UNITED STATES

Ungerstedt, Johanna, Stockholm, SWEDEN

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 2003-369094, filed

on 14 Feb 2003, ABANDONED

PRIORITY INFORMATION: US 2004-577089P 20040604 (60) US 2002-357383P 20020215 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: MINTZ LEVIN COHN FERRIS GLOVSKY & POPEO, 666 THIRD

AVENUE, NEW YORK, NY, 10017, US

NUMBER OF CLAIMS: 27 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Page(s)

LINE COUNT: 2218

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to methods for monitoring patient response to histone deacetylase inhibitors (e.g., suberoylanilide hydroxamic acid (SAHA)) or other therapeutic agents by measuring the level of thioredoxin in body fluids, tissues, and/or cells, such as peripheral blood mononuclear cells, plasma, or serum. The invention also relates to methods of monitoring and/or assisting with the diagnosis of a wide variety of thioredoxin-related diseases and conditions, such as inflammatory diseases, allergic diseases, autoimmune diseases, diseases associated with oxidative stress or diseases characterized by cellular hyperproliferation.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9, Suberoylanilide hydroxamic acid

(use of thioredoxin expression measurements for diagnostics and monitoring treatments with histone deacetylase inhibitors and other therapeutic agents for hyperproliferative diseases)

149647-78-9 USPATFULL RN

Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME) CN

PhNH_C_(CH₂)6_C_NH_OH

ANSWER 9 OF 10 USPATFULL on STN

ACCESSION NUMBER:

2004:166081 USPATFULL Full-text

TITLE:

Method of Treating Leukemia with a Combination of

INVENTOR(S):

Suberoylanilide Hydromaxic Acid and Imatinib Mesylate Bhalla, Kapil N., 12902 Magnolia Drive, MRC-3E 3056D,

Tampa, FL, UNITED STATES 33612

Nimmanapalli, Ramadevi, 12902 Magnolia Drive, MRC-3E

3056D, Tampa, FL, UNITED STATES 33612

PATENT ASSIGNEE(S):

UNIVERSITY OF SOUTH FLORIDA, Tampa; FL, UNITED STATES

(U.S. corporation)

NUMBER KIND DATE US 2004127571 A1 US 2003-605283 A1 PATENT INFORMATION: 20040701 APPLICATION INFO.: 20030919 (10)

> NUMBER DATE ______

PRIORITY INFORMATION:

US 2002-319563P 20020919 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

SMITH & HOPEN PA, 15950 BAY VISTA DRIVE, SUITE 220,

CLEARWATER, FL, 33760

NUMBER OF CLAIMS:

16

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

3 Drawing Page(s)

LINE COUNT:

235

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A method for inducing apoptosis, or increasing the rate or extent of apoptosis, in target cells. The method comprises the steps of contacting the cancer cells with an apoptosis-inducing amount of a tyrosine kinase inhibitor, imatinib mesylate, and a histone deacetylase inhibitor, Suberoylanilide Hydromaxic Acid (SAHA). The method is applicable to ameliorating the resistance of the accelerated and blast phases of CML (CML-BC) to imatinib mesylate.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9, Suberoylanilide hydroxamic acid

(suberoylanilide hydroxamic acid-imatinib mesylate combination for

leukemia treatment)

RN 149647-78-9 USPATFULL

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

L5 ANSWER 10 OF 10 USPATFULL on STN

ACCESSION NUMBER: 2004:166033 U

TITLE:

2004:166033 USPATFULL Full-text

INVENTOR(S):

Methods of treating cancer with HDAC inhibitors
Bacopoulos, Nicholas G., New York, NY, UNITED STATES
Chiao, Judy H., Berkeley Heights, NJ, UNITED STATES

Miller, Thomas A., New York, NY, UNITED STATES

Paradise, Carolyn M., Cortland Manor, NY, UNITED STATES

Richon, Victoria M., Rye, NY, UNITED STATES

NUMBER KIND DATE

PATENT INFORMATION:

US 2004127523 A1 20040701

APPLICATION INFO.:

US 2003-665079 A1 20030916 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2003-379149, filed

on 4 Mar 2003, PENDING

NUMBER DATE

PRIORITY INFORMATION:

US 2002-361759P 20020304 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

MINTZ LEVIN COHN FERRIS GLOVSKY & POPEO, 666 THIRD

AVENUE, NEW YORK, NY, 10017

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

62

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

2792

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to methods of treating cancers, e.g., lymphoma. More specifically, the present invention relates to methods of treating diffuse large B-cell lymphoma (DLBCL), by administration of pharmaceutical compositions comprising HDAC inhibitors, e.g., suberoylanilide hydroxamic acid (SAHA). The oral formulations of the pharmaceutical compositions have favorable pharmacokinetic profiles such as high bioavailability and surprisingly give rise to high blood levels of the active compounds over an extended period of time. The present invention

further provides a safe, daily dosing regimen of these pharmaceutical compositions, which is easy to follow, and which results in a therapeutically effective amount of the HDAC inhibitors in vivo.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9P, SAHA

(histone deacetylase inhibitors for treatment of cancer)

149647-78-9 USPATFULL RN

Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME) CN

PhNH-C-(CH2)6-C-NH-OH

=> d his

L1

L4

(FILE 'HOME' ENTERED AT 12:18:16 ON 01 DEC 2006)

FILE 'REGISTRY' ENTERED AT 12:18:29 ON 01 DEC 2006

STRUCTURE UPLOADED

L2 5 S L1

L3 2 S L1 EXA FULL

FILE 'MEDLINE, CAPLUS, USPATFULL, WPIDS' ENTERED AT 12:19:19 ON 01 DEC

2006

378 S L3

10 S L4 AND "B-CELL LYMPHOMA"

=> s 14 not py>2003

81 L4 NOT PY>2003

=> s 16 and "oral administration"

8 L6 AND "ORAL ADMINISTRATION"

=> d 17 1-8 ibib, abs, hitstr

ANSWER 1 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:405492 CAPLUS Full-text

DOCUMENT NUMBER:

139:115446

TITLE:

Molecular sequelae of histone deacetylase inhibition

in human malignant B cells

AUTHOR(S):

Mitsiades, Nicholas; Mitsiades, Constantine S.; Richardson, Paul G.; McMullan, Ciaran; Poulaki, Vassiliki; Fanourakis, Galinos; Schlossman, Robert; Chauhan, Dharminder; Munshi, Nikhil C.; Hideshima, Teru; Richon, Victoria M.; Marks, Paul A.; Anderson,

Kenneth C.

CORPORATE SOURCE:

Jerome Lipper Multiple Myeloma Center, Department of Medical Oncology, Dana-Farber Cancer Institute,

Harvard Medical School, Boston, MA, USA

SOURCE:

Blood (2003), 101(10), 4055-4062 CODEN: BLOOAW; ISSN: 0006-4971

PUBLISHER:

American Society of Hematology

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Histone acetylation modulates gene expression, cellular differentiation, and survival and is regulated by the opposing activities of histone acetyltransferases (HATs) and histone deacetylases (HDACs). HDAC inhibition results in accumulation of acetylated nucleosomal histones and induces differentiation and/or apoptosis in transformed cells. In this study, we characterized the effect of suberoylanilide hydroxamic acid (SAHA), the prototype of a series of hydroxamic acid-based HDAC inhibitors, in cell lines and patient cells from B-cell malignancies, including multiple myeloma (MM) and related disorders. SAHA induced apoptosis in all tumor cells tested, with increased p21 and p53 protein levels and dephosphorylation of Rb. We also detected cleavage of Bid, suggesting a role for Bcl-2 family members in regulation of SAHA-induced cell death. Transfection of Bcl-2 cDNA into MM.1S cells completely abrogated SAHA-induced apoptosis, confirming its protective role. SAHA did not induce cleavage of caspase-8, -9, or -3 in MM.1S cells during the early phase of apoptosis, and the pan-caspase inhibitor ZVAD-FMK did not protect against SAHA. Conversely, poly(ADP)ribose polymerase (PARP) was cleaved in a pattern indicative of calpain activation, and the calpain inhibitor calpeptin abrogated SAHA-induced cell death. Importantly, SAHA sensitized MM.1S cells to death receptor-mediated apoptosis and inhibited the secretion of interleukin 6 (IL-6) induced in bone marrow stromal cells (BMSCs) by binding of MM cells, suggesting that it can overcome cell adhesion-mediated drug resistance. Our studies delineate the mechanisms whereby HDAC inhibitors mediate anti-MM activity and overcome drug resistance in the BM milieu and provide the framework for clin. evaluation of SAHA, which is bioavailable, well tolerated, and bioactive after oral administration, to improve patient outcome.

IT 149647-78-9, Suberoylanilide hydroxamic acid
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(histone deacetylase inhibitor; authors characterized the effect of
suberoylanilide hydroxamic acid (SAHA) in cell lines and patient cells
from B-cell malignancies, including multiple myeloma (MM) and related
disorders)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:155835 CAPLUS Full-text

DOCUMENT NUMBER:

139:63206

TITLE:

Suberoylanilide hydroxamic acid, a histone deacetylase

inhibitor, ameliorates motor deficits in a mouse model

of Huntington's disease

AUTHOR (S):

Hockly, Emma; Richon, Victoria M.; Woodman, Benjamin;

Smith, Donna L.; Zhou, Xianbo; Rosa, Eddie;

Sathasivam, Kirupa; Ghazi-Noori, Shabnam; Mahal, Amarbirpal; Lowden, Philip A. S.; Steffan, Joan S.; Marsh, J. Lawrence; Thompson, Leslie M.; Lewis,

Cathryn M.; Marks, Paul A.; Bates, Gillian P.

CORPORATE SOURCE:

Medical and Molecular Genetics, Guy's, King's and St. Thomas' School of Medicine, King's College London,

Guy's Hospital, London, SE1 9RT, UK

SOURCE: Proceedings of the National Academy

Proceedings of the National Academy of Sciences of the United States of America (2003), 100(4), 2041-2046

CODEN: PNASA6; ISSN: 0027-8424

PUBLISHER:

National Academy of Sciences

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB Huntington's disease (HD) is an inherited, progressive neurol. disorder that is caused by a CAG/polyglutamine repeat expansion and for which there is no effective therapy. Recent evidence indicates that transcriptional dysregulation may contribute to the mol. pathogenesis of this disease. Supporting this view, administration of histone deacetylase (HDAC) inhibitors has been shown to rescue lethality and photoreceptor neurodegeneration in a Drosophila model of polyglutamine disease. To further explore the therapeutic potential of HDAC inhibitors, we have conducted preclin. trials with suberoylanilide hydroxamic acid (SAHA), a potent HDAC inhibitor, in the R6/2 HD mouse model. We show that SAHA crosses the blood-brain barrier and increases histone acetylation in the brain. We found that SAHA could be administered orally in drinking water when complexed with cyclodextrins. SAHA dramatically improved the motor impairment in R6/2 mice, clearly validating the pursuit of this class of compds. as HD therapeutics.

IT 149647-78-9, Suberoylanilide hydroxamic acid

RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU

(Therapeutic use); BIOL (Biological study); USES (Uses)

(suberoylanilide hydroxamic acid, histone deacetylase inhibitor, ameliorates motor deficits in mouse model of Huntington's disease)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

IT 149647-78-9D, Suberoylanilide hydroxamic acid, complex with cyclodextrin

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (suberoylanilide hydroxamic acid-cyclodextrin complex as an aqueous solution

for oral administration in the treatment of

Huntington's disease)

RN 149647-78-9 CAPLUS

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN ACCESSION NUMBER: 2002:224891 CAPLUS Full-text

DOCUMENT NUMBER:

137:72839

TITLE: The antitumor histone deacetylase inhibitor

suberoylanilide hydroxamic acid exhibits antiinflammatory properties via suppression of

cytokines

AUTHOR (S):

Leoni, Flavio; Zaliani, Andrea; Bertolini, Giorgio; Porro, Giulia; Pagani, Paolo; Pozzi, Pietro; Dona, Giancarlo; Fossati, Gianluca; Sozzani, Silvano; Azam, Tania; Bufler, Philip; Fantuzzi, Giamila; Goncharov, Igor; Kim, Soo-Hyun; Pomerantz, Benjamin J.; Reznikov, Leonid L.; Siegmund, Britta; Dinarello, Charles A.;

Mascagni, Paolo

CORPORATE SOURCE:

Italfarmaco, SpA., Balsamo, 20092, Italy

SOURCE:

Proceedings of the National Academy of Sciences of the

United States of America (2002), 99(5), 2995-3000

CODEN: PNASA6; ISSN: 0027-8424

PUBLISHER:

National Academy of Sciences

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Suberoylanilide hydroxamic acid (SAHA) is a hydroxamic acid-containing hybrid polar mol.; SAHA specifically binds to and inhibits the activity of histone deacetylase. Although SAHA, like other inhibitors of histone deacetylase, exhibits antitumor effects by increasing expression of genes regulating tumor survival, we found that SAHA reduces the production of proinflammatory cytokines in vivo and in vitro. A single oral administration of SAHA to mice dose-dependently reduced circulating TNF-α, IL-1β, IL-6, and IFN-γ induced by lipopolysaccharide (LPS). Administration of SAHA also reduced hepatic cellular injury in mice following i.v. injection of Con A. SAHA inhibited nitric oxide release in mouse macrophages stimulated by the combination of TNF- α plus IFN- γ . Human peripheral blood mononuclear cells stimulated with LPS in the presence of SAHA released less TNF- α , IL-1 β , IL-12, and IFN- γ (50% reduction at 100-200 nM). The production of IFN-y stimulated by IL-18 plus IL-12 was also inhibited by SAHA (85% at 200 nM). However, SAHA did not affect LPS-induced synthesis of the IL-1β precursor, the IL-1 receptor antagonist, or the chemokine IL-8. In addition, IFN-γ induced by anti-CD3 was not suppressed by SAHA. Steady-state mRNA levels for LPS-induced TNF- α and IFN-γ in peripheral blood mononuclear cells were markedly decreased, whereas IL-8 and IL-1 β mRNA levels were unaffected. Because SAHA exhibits antiinflammatory properties in vivo and in vitro, inhibitors of histone deacetylase may stimulate the expression of genes that control the synthesis of cytokines and nitric oxide or hyper-acetylate other targets.

TT 149647-78-9

RN

1.7

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antitumor histone deacetylase inhibitor suberoylanilide hydroxamic acid exhibits antiinflammatory properties via suppression of cytokines) 149647-78-9 CAPLUS

Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME) CN

PhNH-C-(CH2)6-C-NH-OH

REFERENCE COUNT:

37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT ACCESSION NUMBER:

2003:334712 USPATFULL Full-text

TITLE:

Richon, Victoria M., Rye, NY, UNITED STATES INVENTOR(S):

Marks, Paul A., Washington, CT, UNITED STATES Rifkind, Richard A., New York, NY, UNITED STATES

Butler, Lisa M., Athelstone, AUSTRALIA

Method of treating TRX mediated diseases

NUMBER KIND

PATENT INFORMATION:

US 2003235588 A1 US 2003-369094 A1 20031225

APPLICATION INFO.:

20030214 (10)

NUMBER DATE

PRIORITY INFORMATION:

US 2002-357383P 20020215 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HAMILTON, BROOK, SMITH & REYNOLDS, P.C., 530 VIRGINIA

ROAD, P.O. BOX 9133, CONCORD, MA, 01742-9133

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

90

NUMBER OF DRAWINGS:

9 Drawing Page(s)

LINE COUNT:

2095

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides a novel method for treating and/or preventing AΒ thioredoxin (TRX) -mediated diseases and conditions, by administering to a subject in need of such treatment a therapeutically effective amount of a histone deacetylase (HDAC) inhibitor or a pharmaceutically acceptable salt or hydrate thereof. The HDAC inhibitor can alter the expression of a thioredoxin-binding-protein (e.g. TBP-2), which in turn can lead to an altered TRX/thioredoxin-binding-protein cellular binding interaction, resulting in an increase or decrease in the level or activity of cellular TRX, for example the expression level or reducing activity of TRX. Thus the present invention relates to the use of HDAC inhibitors in a method of preventing and/or treating a wide variety of thioredoxin (TRX)-mediated diseases and conditions, such as inflammatory diseases, allergic diseases, autoimmune diseases, diseases associated with oxidative stress or diseases characterized by cellular hyperproliferation.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9, SAHA

(use of histone deacetylase inhibitors for preventing/treating thioredoxin (TRX) mediated diseases or conditions associated with inflammation and cellular hyperproliferation)

RN149647-78-9 USPATFULL

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

PhNH_C-(CH₂)6-C-NH-OH

ANSWER 5 OF 8 USPATFULL on STN

ACCESSION NUMBER:

2003:166667 USPATFULL Full-text

TITLE: Method of treating autoimmune diseases

INVENTOR(S): Kammer, Gary M., Lewisville, NC, UNITED STATES Mishra, Nilamadhab, Winston-Salem, NC, UNITED STATES

KIND DATE NUMBER _______

PATENT INFORMATION:

APPLICATION INFO.:

US 2003114525 A1 20030619 US 2002-151481 A1 20020520 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2000-718195, filed

on 20 Nov 2000, ABANDONED

NUMBER ______

PRIORITY INFORMATION:

WO 2001-US43871 20011119

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE: MYERS BIGEL SIBLEY & SAJOVEC, PO BOX 37428, RALEIGH,

NC, 27627

NUMBER OF CLAIMS:

36

EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

25 Drawing Page(s)

LINE COUNT:

1464

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB

A method of treating an autoimmune disease comprising administering to the subject a treatment effective amount of a histone hyperacetylating agent, or

a pharmaceutically acceptable salt thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9, Suberoylanilide Hydroxamic acid

(method of treating autoimmune diseases using a histone

hyperacetylating agent)

149647-78-9 USPATFULL RN

Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME) CN

PhNH_C_ (CH2) 6 _ C_NH_OH

L7 ANSWER 6 OF 8 USPATFULL on STN

ACCESSION NUMBER:

INVENTOR(S):

2003:140935 USPATFULL Full-text

TITLE:

Modulation of gene expression by combination therapy

Besterman, Jeffrey M., Bai D'Urfe, CANADA

MacLeod, Robert A., Westmount, CANADA

Siders, William M., Watertown, MA, UNITED STATES

Li, Zuomei, Kirkland, CANADA

NUMBER KIND DATE ______ US 2003096777 A1 20030522

PATENT INFORMATION: APPLICATION INFO.:

A1 20020514 (10)

RELATED APPLN. INFO.:

US 2002-145493 Division of Ser. No. US 1999-420692, filed on 19 Oct

1999, PENDING

NUMBER DATE

PRIORITY INFORMATION:

US 1998-104804P 19981019 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

KEOWN & ASSOCIATES, 500 WEST CUMMINGS PARK, SUITE 1200,

WOBURN, MA, 01801

NUMBER OF CLAIMS:

50

EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

31 Drawing Page(s)

LINE COUNT:

2704

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB

The invention relates to the modulation of gene expression. In particular, the invention relates to compositions comprising antisense oligonucleotides which inhibit expression of a gene in operable association with protein effectors of a product of that gene, and methods of using the same.

In addition, the invention relates to the modulation of mammalian gene expression regulated by methylation.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9

(antisense oligonucleotide and gene product protein effector for gene expression modulation)

RN 149647-78-9 USPATFULL

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

PhNH_C_(CH₂)6_C_NH_OH

L7 ANSWER 7 OF 8 USPATFULL on STN

ACCESSION NUMBER:

2003:120185 USPATFULL Full-text

TITLE:

Method of treating autoimmune diseases

INVENTOR(S):

Kammer, Gary M., Lewisville, NC, UNITED STATES

Mishra, Nilamadhab, Winston-Salem, NC, UNITED STATES

NUMBER KIND DATE

PATENT INFORMATION:

US 2003082666 A1 20030501

APPLICATION INFO.:

US 2002-187586 A1 20020702 (10)

RELATED APPLN. INFO.:

Division of Ser. No. US 2000-718195, filed on 20 Nov

2000, ABANDONED

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

MYERS BIGEL SIBLEY & SAJOVEC, PO BOX 37428, RALEIGH,

NC, 27627

NUMBER OF CLAIMS:

29

EXEMPLARY CLAIM:

-

NUMBER OF DRAWINGS:

3 Drawing Page(s)

LINE COUNT:

843

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB

A method of treating an autoimmune disease (for example, Systemic Lupus Erythematosus) comprises administering to the subject a treatment effective amount of a histone hyperacetylating agent, or a pharmaceutically acceptable salt thereof. Methods of screening compounds useful for the treatment of autoimmune disease are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9

(method of treating autoimmune diseases with histone hyperacetylating agent)

RN 149647-78-9 USPATFULL

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

PhNH-C-(CH₂)6-C-NH-OH

L7 ANSWER 8 OF 8 USPATFULL on STN

ACCESSION NUMBER:

2002:337369 USPATFULL Full-text

TITLE:

MAP-2 as a determinant of metastatic potential Setaluri, Vijayasaradhi, Winston-Salem, NC, UNITED

INVENTOR(S): Setalu: STATES

Fang, Dong, Winston-Salem, NC, UNITED STATES White, Wain, Winston-Salem, NC, UNITED STATES

•	NUMBER	KIND	DATE	•	
					•
PATENT INFORMATION:	US 2002192727	A1	20021219		•
	US 6613534	B2	20030902		
APPLICATION INFO.:	US 2001-812348	A1	20010320	(9)	
DOCUMENT TYPE:	Utility			•	
FILE SEGMENT:	APPLICATION				
LEGAL REPRESENTATIVE:	KILPATRICK STOCK	TON LLP	, 1001 WES'	T FOURTH STREET,	,
	WINSTON-SALEM, · N	C, 2710	1		
NUMBER OF CLAIMS:	68				
EXEMPLARY CLAIM:	1	•			
NUMBER OF DRAWINGS:	8 Drawing Page(s)			
T TMD COIDIN	1720				

LINE COUNT: 1730

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to detection of MAP-2 (microtubule associated protein-2) as a marker to determine the metastatic potential of a tumor, including tumors derived from the neural crest such as melanomas, gliomas, Schwanomas, chromocytomas and small cell lung cancer. In one aspect, the invention comprises a method for determining the metastatic potential of a tumor sample, wherein decreased levels of MAP-2 expression in a test sample relative to controls indicates that the sample has increased metastatic potential as compared to the control. In another aspect, the invention comprises a method to prevent tumor progression in metastatic melanoma by increasing levels of MAP-2 protein in cells.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149647-78-9

(microtubule associated protein-2 (MAP-2) and its role as an indicator of metastatic potential)

RN 149647-78-9 USPATFULL

CN Octanediamide, N-hydroxy-N'-phenyl- (9CI) (CA INDEX NAME)

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(FILE 'HOME' ENTERED AT 12:18:16 ON 01 DEC 2006)

FILE 'REGISTRY' ENTERED AT 12:18:29 ON 01 DEC 2006

L1 STRUCTURE UPLOADED

L2 5 S L1

L3 2 S L1 EXA FULL

FILE 'MEDLINE, CAPLUS, USPATFULL, WPIDS' ENTERED AT 12:19:19 ON 01 DEC 2006

L4 378 S L3

L5 10 S L4 AND "B-CELL LYMPHOMA"

L6 81 S L4 NOT PY>2003

L7 8 S L6 AND "ORAL ADMINISTRATION"

=> s 16 and cancer

L8 36 L6 AND CANCER

=> s 16 and lymphoma

L9 2 L6 AND LYMPHOMA

=> d 19 1-2 ibib, abs

L9 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN ACCESSION NUMBER: 2003:847686 CAPLUS Full-text

DOCUMENT NUMBER: 139:358271

TITLE: Phase I clinical trial of histone deacetylase

inhibitor: suberoylanilide hydroxamic acid

administered intravenously

AUTHOR(S): Kelly, Wm. Kevin; Richon, Victoria M.; O'Connor, Owen;

Curley, Tracy; MacGregor-Curtelli, Barbara; Tong, William; Klang, Mark; Schwartz, Lawrence; Richardson, Stacie; Rosa, Eddie; Drobnjak, Marija; Cordon-Cordo, Carlos; Chiao, Judy H.; Rifkind, Richard; Marks, Paul

A.; Scher, Howard

CORPORATE SOURCE: Genitourinary Oncology Service, Department of

Medicine, Memorial Sloan-Kettering Cancer Center and Joan and Sanford Weill Medical College of Cornell

University, New York, NY, 10021, USA

SOURCE: Clinical Cancer Research (2003), 9(10, Pt. 1),

3578-3588

CODEN: CCREF4; ISSN: 1078-0432

PUBLISHER: American Association for Cancer Research

DOCUMENT TYPE: Journal LANGUAGE: English

AB To evaluate the safety, pharmacokinetics, and biol. activity of suberoylanilide hydroxamic acid (SAHA) administered by 2-h i.v. infusion in patients with advanced cancer. SAHA was administered for 3 days every 21 days in part A and 5 days for 1-3 wk in part B. Dose escalation proceeded independently in patients with solid tumor and hematol. malignancies (part B only). Pharmacokinetic studies were performed along with assessment of acetylated histones in peripheral blood mononuclear cells and tumor tissues. No dose-limiting toxicities were observed in 8 patients enrolled in part A

(75, 150, 300, 600, and 900 mg/m2/day). Among 12 hematol. and 17 solid tumor patients enrolled in part B (300, 600, and 900 mg/m2/day), therapy was delayed ≥1 wk for grade 3/4 leukopenia and/or thrombocytopenia in 2 of 5 hematol. patients at 600 mg/m2/day + 5 days for 3 wk. The maximal-tolerated dose was 300 mg/m2/day + 5 days for 3 wk for hematol. patients. One solid patient on 900 mg/m2/day + 5 days for 3 wk developed acute respiratory distress and grade 3 hypotension. The cohort was expanded to 6 patients, and no addnl. doselimiting toxicities were observed Mean terminal half-life ranged from 21 to 58 min, and there, was dose-proportional increase in area under the curve. An accumulation of acetylated histones in peripheral blood mononuclear cells up to 4 h postinfusion was observed at higher dose levels. Posttherapy tumor biopsies showed an accumulation of acetylated histones by immunohistochem. Four (2 lymphoma and 2 bladder) patients had objective tumor regression with clin. improvement in tumor related symptoms. Daily i.v. SAHA is well tolerated, inhibits the biol. target in vivo, and has antitumor activity in solid and hematol. tumors.

REFERENCE COUNT:

17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 2 USPATFULL on STN

ACCESSION NUMBER:

2002:323219 USPATFULL Full-text

TITLE:

Use of retinoids plus histone deacetylase inhibitors to

inhibit the growth of solid tumors

INVENTOR(S):

Gudas, Lorraine J., New York, NY, UNITED STATES Nanus, David, New Rochelle, NY, UNITED STATES

PATENT INFORMATION: APPLICATION INFO.:

US 2002183388 A1 20021205 US 2002-61101 A1 20020201 (10)

NUMBER DATE

PRIORITY INFORMATION:

US 2001-265651P 20010201 (60)

DOCUMENT TYPE: FILE SEGMENT: Utility APPLICATION

LEGAL REPRESENTATIVE:

Michael L. Goldman, Esq., NIXON PEABODY LLP, Clinton

Square, P.O. Box 31051, Rochester, NY, 14603-1051

NUMBER OF CLAIMS: 11

EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 20 Drawing Page(s)

549

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides a method of inhibiting growth of solid tumors in an animal which comprises administering an effective amount of trichostatin A (TSA) to an animal in need of such treatment. The present invention also provides a method of inhibiting growth of solid tumors in an animal which comprises administering an effective amount of a histone deacetylase inhibitor and a retinoid to an animal in need of such treatment. Examples of solid tumors which may be treated using the methods of the invention include but are not limited to carcinomas of the head and neck, breast, skin, kidney, oral cavity, colon, prostate, pancreas and lung.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> d his

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FILE 'REGISTRY' ENTERED AT 12:18:29 ON 01 DEC 2006
                STRUCTURE UPLOADED
Ll
              5 S L1
L2
L3
              2 S L1 EXA FULL
     FILE 'MEDLINE, CAPLUS, USPATFULL, WPIDS' ENTERED AT 12:19:19 ON 01 DEC
            378 S L3
L4
L5
             10 S L4 AND "B-CELL LYMPHOMA"
L6
             81 S L4 NOT PY>2003
             8 S L6 AND "ORAL ADMINISTRATION"
L7
             36 S L6 AND CANCER
L8
             2 S L6 AND LYMPHOMA
L9
=> s 16 and "orally"
             7 L6 AND "ORALLY"
=> d 110 1-7 ibib, abs
L10 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN
                         2003:155835 CAPLUS Full-text
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         139:63206
                         Suberoylanilide hydroxamic acid, a histone deacetylase
TITLE:
                         inhibitor, ameliorates motor deficits in a mouse model
                         of Huntington's disease
                         Hockly, Emma; Richon, Victoria M.; Woodman, Benjamin;
AUTHOR (S):
                         Smith, Donna L.; Zhou, Xianbo; Rosa, Eddie;
                         Sathasivam, Kirupa; Ghazi-Noori, Shabnam; Mahal,
                         Amarbirpal; Lowden, Philip A. S.; Steffan, Joan S.;
                         Marsh, J. Lawrence; Thompson, Leslie M.; Lewis,
                         Cathryn M.; Marks, Paul A.; Bates, Gillian P.
                         Medical and Molecular Genetics, Guy's, King's and St.
CORPORATE SOURCE:
                         Thomas' School of Medicine, King's College London,
                         Guy's Hospital, London, SE1 9RT, UK
                         Proceedings of the National Academy of Sciences of the
SOURCE:
                         United States of America (2003), 100(4), 2041-2046
                         CODEN: PNASA6; ISSN: 0027-8424
PUBLISHER:
                         National Academy of Sciences
DOCUMENT TYPE:
                         Journal
LANGUAGE:
                         English
     Huntington's disease (HD) is an inherited, progressive neurol. disorder that
AR
     is caused by a CAG/polyglutamine repeat expansion and for which there is no
     effective therapy. Recent evidence indicates that transcriptional
     dysregulation may contribute to the mol. pathogenesis of this disease.
     Supporting this view, administration of histone deacetylase (HDAC) inhibitors
     has been shown to rescue lethality and photoreceptor neurodegeneration in a
     Drosophila model of polyglutamine disease. To further explore the therapeutic
     potential of HDAC inhibitors, we have conducted preclin. trials with
     suberoylanilide hydroxamic acid (SAHA), a potent HDAC inhibitor, in the R6/2
     HD mouse model. We show that SAHA crosses the blood-brain barrier and
     increases histone acetylation in the brain. We found that SAHA could be
     administered orally in drinking water when complexed with cyclodextrins. SAHA
     dramatically improved the motor impairment in R6/2 mice, clearly validating
     the pursuit of this class of compds. as HD therapeutics.
REFERENCE COUNT:
                         36
                               THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
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L10 ANSWER 2 OF 7 USPATFULL on STN
ACCESSION NUMBER: 2003:334712 USPATFULL Full-text

TITLE: INVENTOR(S):

Method of treating TRX mediated diseases Richon, Victoria M., Rye, NY, UNITED STATES Marks, Paul A., Washington, CT, UNITED STATES Rifkind, Richard A., New York, NY, UNITED STATES

Butler, Lisa M., Athelstone, AUSTRALIA

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.: US 2003235588 A1 20031225 US 2003-369094 A1 20030214 (10)

NUMBER DATE _ -----

PRIORITY INFORMATION:

US 2002-357383P 20020215 (60)

DOCUMENT TYPE: Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE: HAMILTON, BROOK, SMITH & REYNOLDS, P.C., 530 VIRGINIA

ROAD, P.O. BOX 9133, CONCORD, MA, 01742-9133

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

9 Drawing Page(s)

LINE COUNT:

2095

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides a novel method for treating and/or preventing thioredoxin (TRX)-mediated diseases and conditions, by administering to a subject in need of such treatment a therapeutically effective amount of a histone deacetylase (HDAC) inhibitor or a pharmaceutically acceptable salt or hydrate thereof. The HDAC inhibitor can alter the expression of a thioredoxin-binding-protein (e.g. TBP-2), which in turn can lead to an altered TRX/thioredoxin-binding-protein cellular binding interaction, resulting in an increase or decrease in the level or activity of cellular TRX, for example the expression level or reducing activity of TRX. Thus the present invention relates to the use of HDAC inhibitors in a method of preventing and/or treating a wide variety of thioredoxin (TRX)-mediated diseases and conditions, such as inflammatory diseases, allergic diseases, autoimmune diseases, diseases associated with oxidative stress or diseases characterized by cellular hyperproliferation.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 3 OF 7 USPATFULL on STN

ACCESSION NUMBER:

2002:323219 USPATFULL Full-text

TITLE:

Use of retinoids plus histone deacetylase inhibitors to

inhibit the growth of solid tumors

INVENTOR(S):

Gudas, Lorraine J., New York, NY, UNITED STATES Nanus, David, New Rochelle, NY, UNITED STATES

NUMBER KIND DATE ------US 2002183388 A1 20021205

PATENT INFORMATION:

APPLICATION INFO.:

US 2002-61101 A1 20020201 (10)

NUMBER DATE -----

PRIORITY INFORMATION:

US 2001-265651P 20010201 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Michael L. Goldman, Esq., NIXON PEABODY LLP, Clinton Square, P.O. Box 31051, Rochester, NY, 14603-1051

NUMBER OF CLAIMS: 11 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 20 Drawing Page(s)

LINE COUNT: 549

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides a method of inhibiting growth of solid tumors in an animal which comprises administering an effective amount of trichostatin A (TSA) to an animal in need of such treatment. The present invention also provides a method of inhibiting growth of solid tumors in an animal which comprises administering an effective amount of a histone deacetylase inhibitor and a retinoid to an animal in need of such treatment. Examples of solid tumors which may be treated using the methods of the invention include but are not limited to carcinomas of the head and neck, breast, skin, kidney, oral cavity, colon, prostate, pancreas and lung.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 4 OF 7 USPATFULL on STN

ACCESSION NUMBER: 2000:88196 USPATFULL Full-text

TITLE: Potent inducers of terminal differentiation and methods

of use thereof

INVENTOR(S): Breslow, Ronald, Englewood, NJ, United States

Marks, Paul A., Bridgewater, CT, United States Rifkind, Richard A., New York, NY, United States Jursic, Branko, New Orleans, LA, United States

PATENT ASSIGNEE(S): Sloan-Kettering Institute for Cancer Research, New

York, NY, United States (U.S. corporation)

The Trustees of Columbia University in the City of New York, New York, NY, United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6087367 20000711
APPLICATION INFO.: US 1999-314195 19990518 (9)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1994-222685, filed on 4 Apr

1994 which is a division of Ser. No. US 1991-771760, filed on 4 Oct 1991, now patented, Pat. No. US 5369108,

issued on 29 Nov 1994

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Kight, John

ASSISTANT EXAMINER: Covington, Raymond

LEGAL REPRESENTATIVE: White, John P.Cooper & Dunham LLP

NUMBER OF CLAIMS: 19
EXEMPLARY CLAIM: 1
LINE COUNT: 1379

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides the compound having the structure: ##STR1## wherein each of R.sub.1 and R.sub.2 are independently the same as or different from each other; when R.sub.1 and R.sub.2 are the same, each is a substituted or unsubstituted arylamino, cycloalkylamino, pyridineamino, piperidino, 9-purine-6-amine, or thiozoleamino group; when R.sub.1 and R.sub.2 are different, R.sub.1 =R.sub.3 --N--R.sub.4, wherein each of R.sub.3 and R.sub.4 are independently the same as or different from each other and are a hydrogen atom, a hydroxyl group, a substituted or unsubstituted, branched or unbranched alkyl, alkenyl, cycloalkyl, aryl, alkyloxy, aryloxy, arylalkyloxy, or pyridine group, or R.sub.3 and R.sub.4 bond together to form a piperidine group and R.sub.2 is a hydroxylamino,

hydroxyl, amino, alkylamino, dialkylamino or alkyloxy group; and n is an integer from about 4 to about 8.

The present invention also provides a method of selectively inducing terminal differentiation of neoplastic cells and thereby inhibiting proliferation of such cells. Moreover, the present invention provides a method of treating a patient having a tumor characterized by proliferation of neoplastic cells. Lastly, the present invention provides a pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically acceptable amount of the compound above.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 5 OF 7 USPATFULL on STN

ACCESSION NUMBER: 1999:89192 USPATFULL Full-text

TITLE: Potent inducers of terminal differentiation and methods

of use thereof

INVENTOR(S): Breslow, Ronald, Englewood, NJ, United States

Marks, Paul A., Bridgewater, CT, United States Rifkind, Richard A., New York, NY, United States Jursic, Branko, New Orleans, LA, United States

PATENT ASSIGNEE(S): Sloan-Kettering Institute for Cancer Research, New

York, NY, United States (U.S. corporation)

The Trustees of Columbia University in the City of New York, New York, NY, United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5932616 19990803 APPLICATION INFO.: US 1994-222685 19940404 (8)

RELATED APPLN. INFO.: Division of Ser. No. US 1991-771760, filed on 4 Oct

1991, now patented, Pat. No. US 5369108

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Kight, John

ASSISTANT EXAMINER: Covington, Raymond

LEGAL REPRESENTATIVE: White, John P.Cooper & Dunham LLP

NUMBER OF CLAIMS: 27
EXEMPLARY CLAIM: 1
LINE COUNT: 1418

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides the compound having the structure: ##STR1## wherein each of R.sub.1 and R.sub.2 are independently the same as or different from each other; when R.sub.1 and R.sub.2 are the same, each is a substituted or unsubstituted arylamino, cycloalkylamino, pyridineamino, piperidino, 9-purine-6-amine, or thiozoleamino group; when R.sub.1 and R.sub.2 are different, R.sub.1 =R.sub.3 --N--R.sub.4, wherein each of R.sub.3 and R.sub.4 are independently the same as or different from each other and are a hydrogen atom, a hydroxyl group, a substituted or unsubstituted, branched or unbranched alkyl, alkenyl, cycloalkyl, aryl, alkyloxy, aryloxy, arylalkyloxy, or pyridine group, or R.sub.3 and R.sub.4 bond together to form a piperidine group and R.sub.2 is a hydroxylamino, hydroxyl, amino, alkylamino, dialkylamino or alkyloxy group; and n is an integer from about 4 to about 8.

The present invention also provides a method of selectively inducing terminal differentiation of neoplastic cells and thereby inhibiting

proliferation of such cells. Moreover, the present invention provides a method of treating a patient having a tumor characterized by proliferation of neoplastic cells. Lastly, the present invention provides a pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically acceptable amount of the compound above.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 6 OF 7 USPATFULL on STN

ACCESSION NUMBER: 97:120628 USPATFULL Full-text

TITLE: Potent inducers of terminal differentiation and method

of use thereof

Breslow, Ronald, Englewood, NJ, United States INVENTOR(S):

Marks, Paul A., Washington, CT, United States Rifkind, Richard A., New York, NY, United States

PATENT ASSIGNEE(S): Sloan-Kettering Institute for Cancer Research, New

York, NY, United States (U.S. corporation)

KIND DATE NUMBER US 5700811 19971223

PATENT INFORMATION: APPLICATION INFO.:

US 1994-246363 19940519 (8)

Continuation-in-part of Ser. No. US 1991-771760, filed RELATED APPLN. INFO.:

on 4 Oct 1991, now patented, Pat. No. US 5369108

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Ivy, C. Warren PRIMARY EXAMINER: Covington, Raymond ASSISTANT EXAMINER:

LEGAL REPRESENTATIVE:

White, John P. NUMBER OF CLAIMS: 18

EXEMPLARY CLAIM: LINE COUNT: 1585

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention is directed to compounds having the structure: ##STR1## AB wherein R.sub.1 and R.sub.2 are independently the same as or different from each other; when R.sub.1 and R.sub.2 are the same, each is a substituted or unsubstituted arylamino, cycloalkylamino, pyridineamino, piperidino, 9purine-6-amine or thiazoleamino group; when R. sub.1 and R. sub.2 are different, R.sub.1 = R.sub.3 -- N-- R.sub.4, and n is an integer from about 4 to about 8. This invention also provides a method of selectively inducing terminal differentiation of neoplastic cells and thereby inhibiting proliferation of such cells. The invention further provides pharmaceutical compositions comprising a therapeutically effective amount of the compounds of the present invention and a pharmaceutically acceptable carrier.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 7 OF 7 USPATFULL on STN

94:104578 USPATFULL Full-text ACCESSION NUMBER:

TITLE: Potent inducers of terminal differentiation and methods

of use thereof

Breslow, Ronald, Englewood, NJ, United States INVENTOR(S):

> Marks, Paul A., Bridgewater, CT, United States Rifkind, Richard A., New York, NY, United States Jursic, Branko, New Orleans, LA, United States Sloan-Kettering Institute for Cancer Research, New

PATENT ASSIGNEE(S):

York, NY, United States (U.S. corporation)

The Trustees of Columbia, New York, NY, United States

(U.S. corporation)

•	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 5369108		19941129	
APPLICATION INFO.:	US 1991-771760		19911004	(7)
DOCUMENT TYPE:	Utility			
FILE SEGMENT:	Granted			
PRIMARY EXAMINER:	Ivy, C. Warren			

PRIMARY EXAMINER: Ivy, C. Warren
ASSISTANT EXAMINER: Covington, Raymond
LEGAL REPRESENTATIVE: White, John P.

NUMBER OF CLAIMS: 17
EXEMPLARY CLAIM: 1
LINE COUNT: 1291

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides the compound having the structure: ##STR1## wherein each of R.sub.1 and R.sub.2 are independently the same as or different from each other; when R.sub.1 and R.sub.2 are the same, each is a substituted or unsubstituted arylamino, cycloalkylamino, pyridineamino, piperidino, 9-purine-6-amine, or thiozoleamino group; when R.sub.1 and R.sub.2 are different, R.sub.1 =R.sub.3 --N--R.sub.4, wherein each of R.sub.3 and R.sub.4 are independently the same as or different from each other and are a hydrogen atom, a hydroxyl group, a substituted or unsubstituted, branched or unbranched alkyl, alkenyl, cycloalkyl, aryl, alkyloxy, aryloxy, arylalkyloxy, or pyridine group, or R.sub.3 and R.sub.4 bond together to form a piperidine group and R.sub.2 is a hydroxylamino, hydroxyl, amino, alkylamino, dialkylamino or alkyloxy group; and n is an integer from about 4 to about 8.

The present invention also provides a method of selectively inducing terminal differentiation of neoplastic cells and thereby inhibiting proliferation of such cells. Moreover, the present invention provides a method of treating a patient having a tumor characterized by proliferation of neoplastic cells. Lastly, the present invention provides a pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically acceptable amount of the compound above.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> d his

(FILE 'HOME' ENTERED AT 12:18:16 ON 01 DEC 2006)

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FILE 'REGISTRY' ENTERED AT 12:18:29 ON 01 DEC 2006
L1 STRUCTURE UPLOADED
L2 5 S L1
L3 2 S L1 EXA FULL
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FILE 'MEDLINE, CAPLUS, USPATFULL, WPIDS' ENTERED AT 12:19:19 ON 01 DEC 2006

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L4 378 S L3
L5 10 S L4 AND "B-CELL LYMPHOMA"
L6 81 S L4 NOT PY>2003
L7 8 S L6 AND "ORAL ADMINISTRATION"
L8 36 S L6 AND CANCER
L9 2 S L6 AND LYMPHOMA
L10 7 S L6 AND "ORALLY"
```

---Logging off of STN---